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REMARKS

Applicants enclose ten replacement sheets of formal drawings to comply with 37 C.F.R. 1.84 and 37 C.F.R. 1.121, each labeled as a “replacement sheet.” These drawings replace those originally filed informal drawings in this application. The replacement sheets reflect the amendments to the originally filed informal drawings as discussed previously. Support for these amendments is recited below. Applicants have also provided marked-up versions of the relevant sheets of originally filed informal drawings indicating these amendments, each labeled as a “marked-up version.” Applicants further submit that these amendments add no new matter.

Additionally, Applicants have amended the specification to incorporate the text presented in the original drawings. Applicants recite support for these amendments below and submit that they add no new matter.

Amendments to the Drawings

Support for amendments to the drawings can be found in the drawings as originally filed.

Amendments to the Specification

Support for amendments to paragraph 16 can be found in Figures 1A and 1B on sheet 1, and in Figure 1C on sheet 2 of the drawings as originally filed.

Support for amendments to paragraph 17 can be found in Figure 2A on sheet 3, in Figure 2B on sheet 4, and in Figure 2C on sheet 5 of the drawings as originally filed.

Support for amendments to paragraph 18 can be found in Figure 3A on sheet 6, and in Figure 3B on sheet 7 of the drawings as originally filed.

Support for amendments to paragraph 19 can be found in Figure 4 on sheet 8 of the drawings as originally filed.

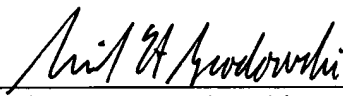
CONCLUSION

Applicants respectfully request entry of this Preliminary Amendment before examination of the above-identified patent application. Applicants believe that the foregoing amendments introduce no new matter. Early favorable action is respectfully solicited.

Respectfully submitted,

Date: July 21, 2004
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Limited Effectiveness of Repeated Delivery of IL18 Adenoviral Vector

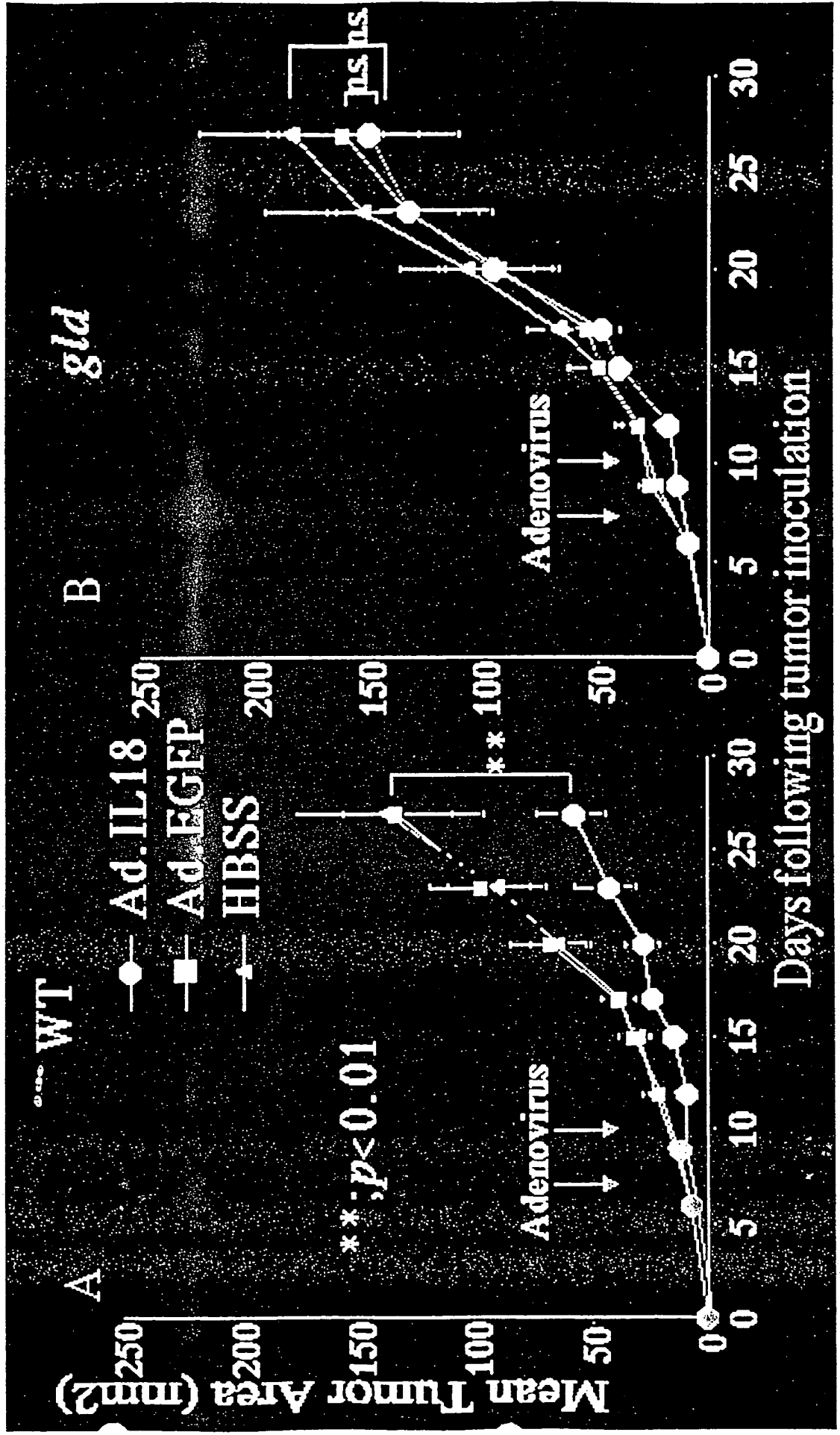


Figure 1A and 1B

FasL Dependent Induction of Cytotoxicity

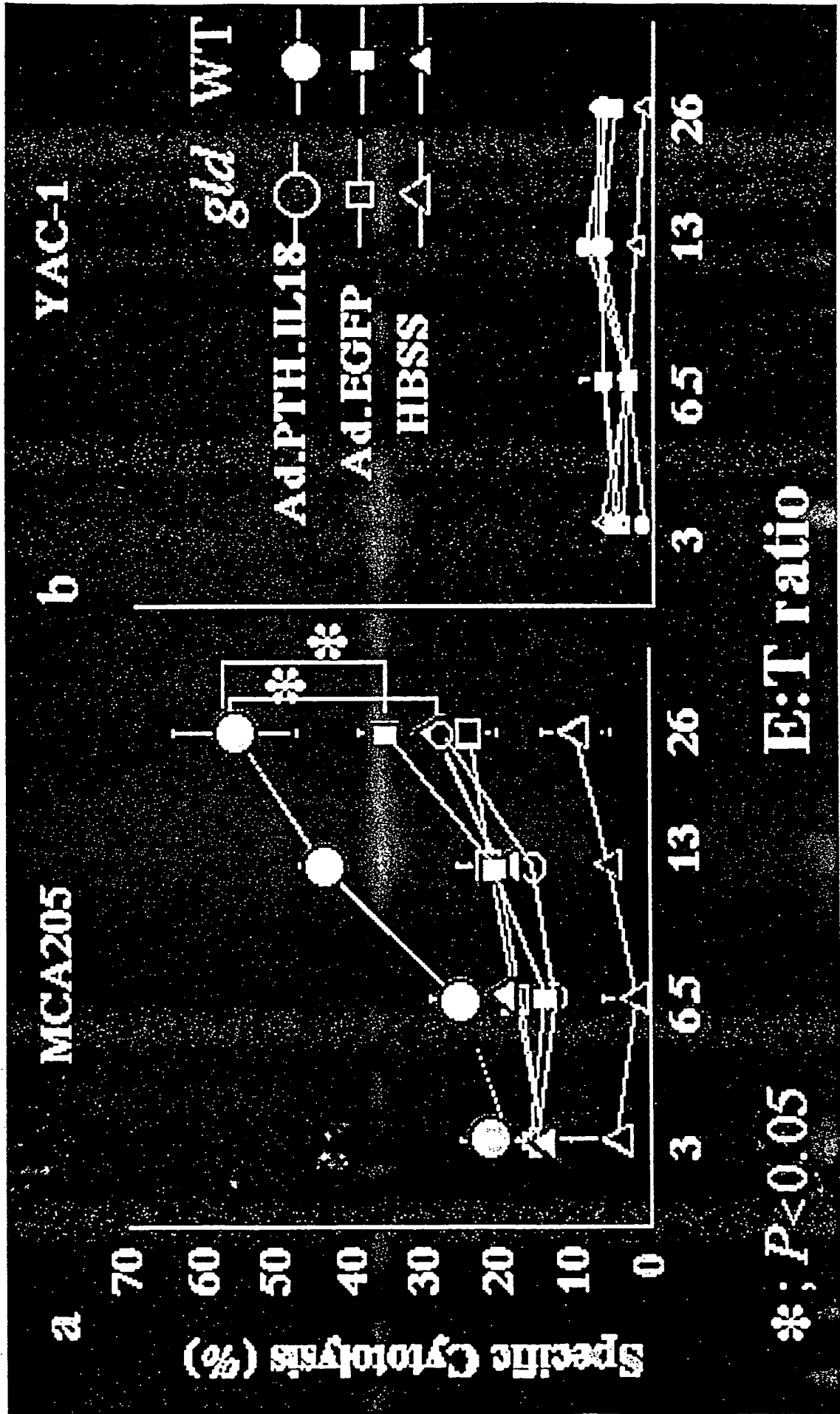


Figure 1c.

DC Combination With Adenoviral Delivery of IL18 Mediates Profound Antitumor Activity in MCA 205

Injected MCA205 tumors

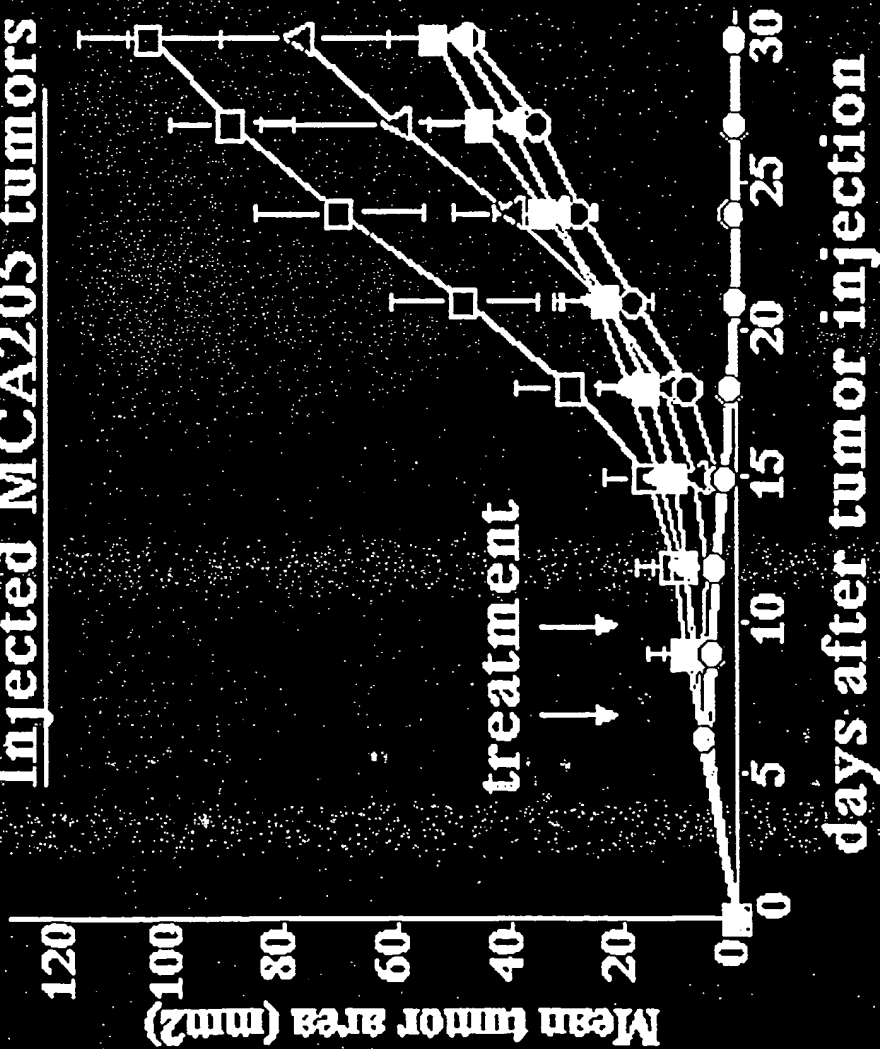
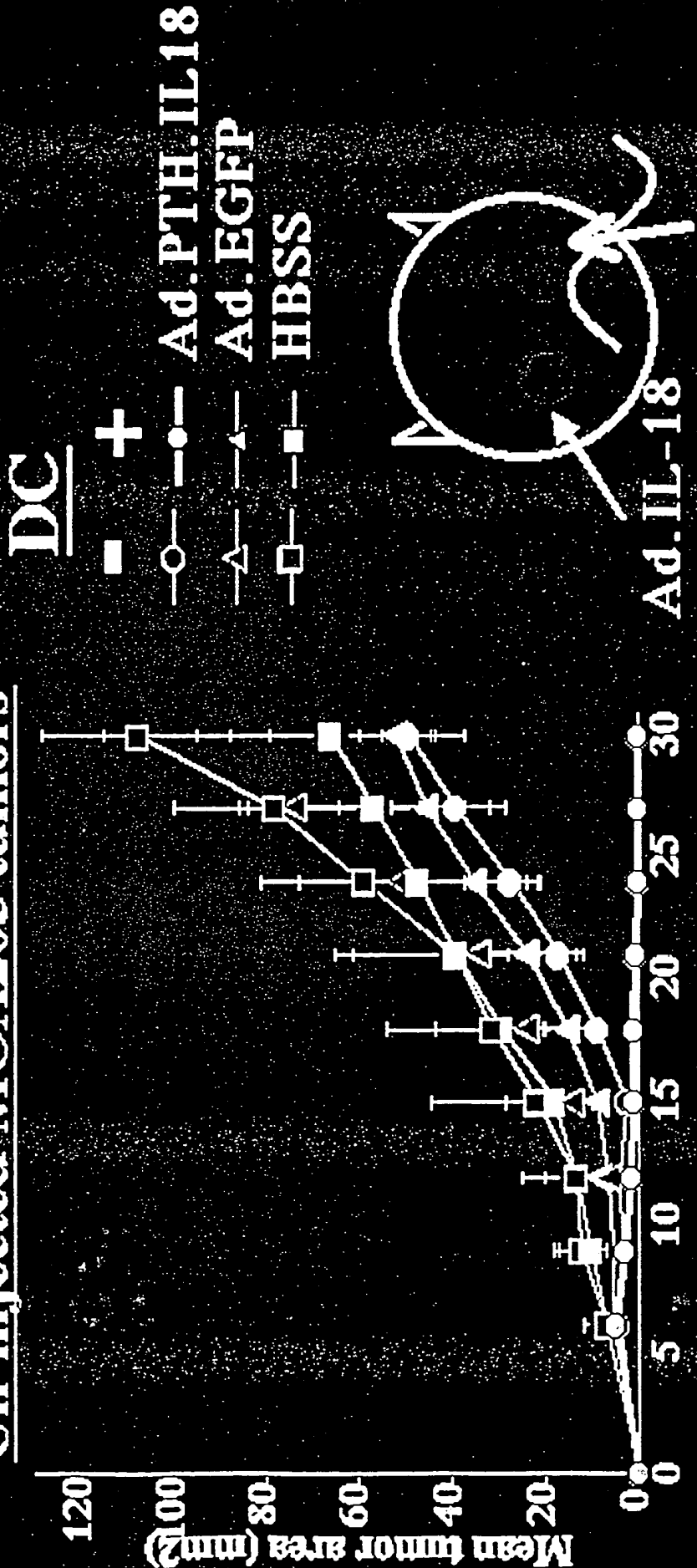


Figure 2A

Interleukin 18 Mediates Distant Antitumor Effects When Coupled with DC

Un-injected MCA205 tumors



days after tumor injection DC Un-injected tumor
 Similar findings with MC38 Colorectal Adenocarcinoma

Figure 2B

Development of Systemic Immunity with IL18 and Dendritic Cell Coadministration

Effector:

Lymph Node Cells

Splenocytes

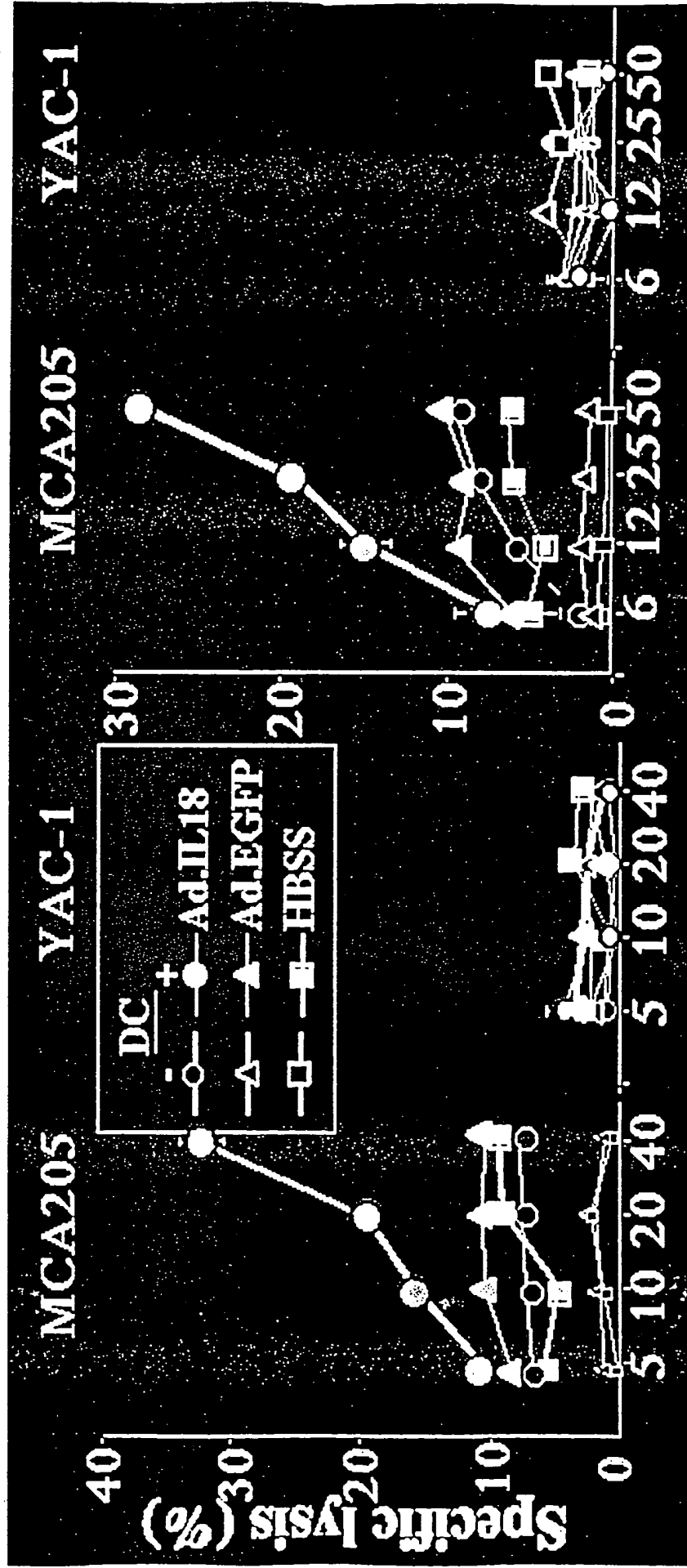


Figure 2C

Fine Specificity of the Antitumor Cytolytic Response Mediated by IL18 and DCs

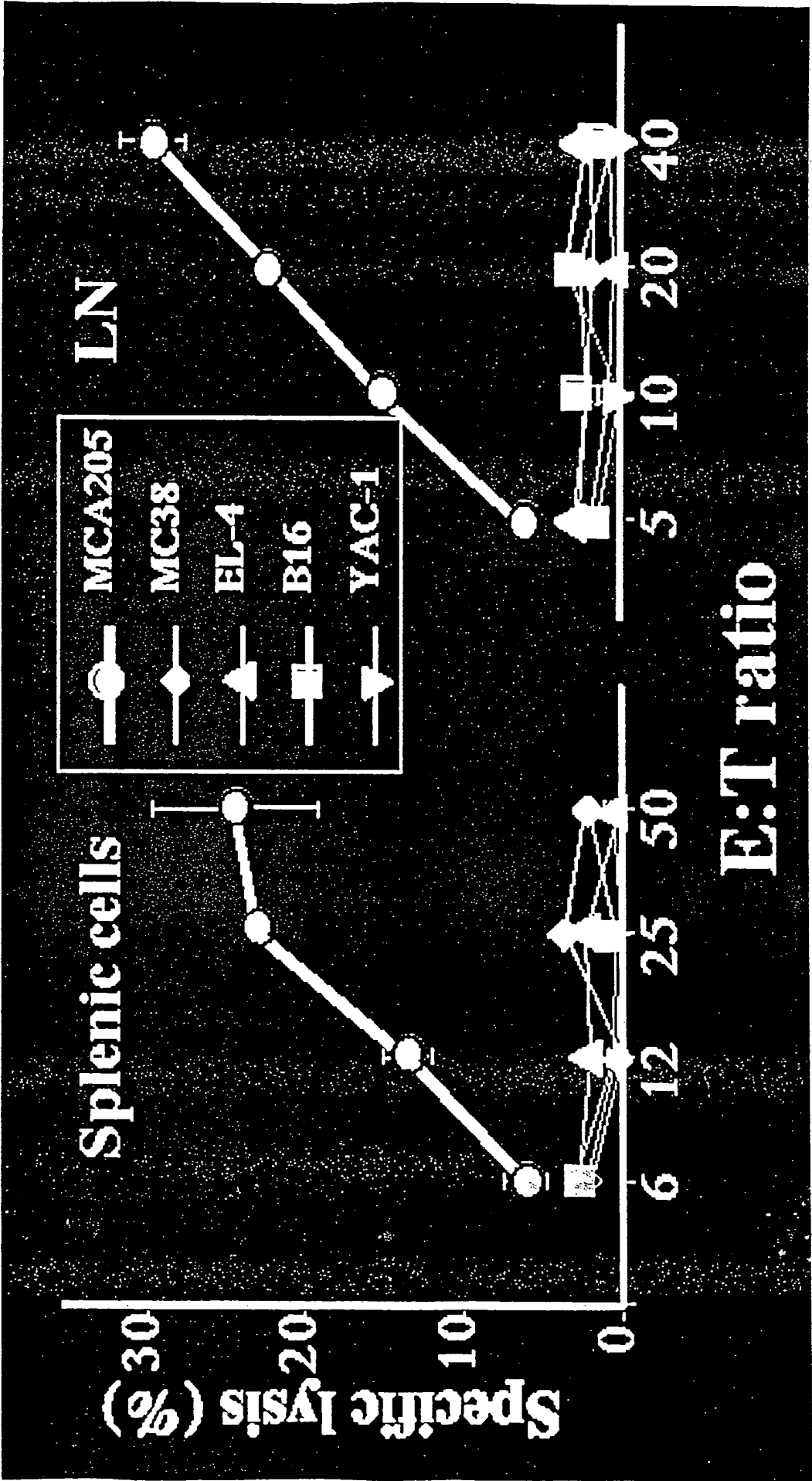
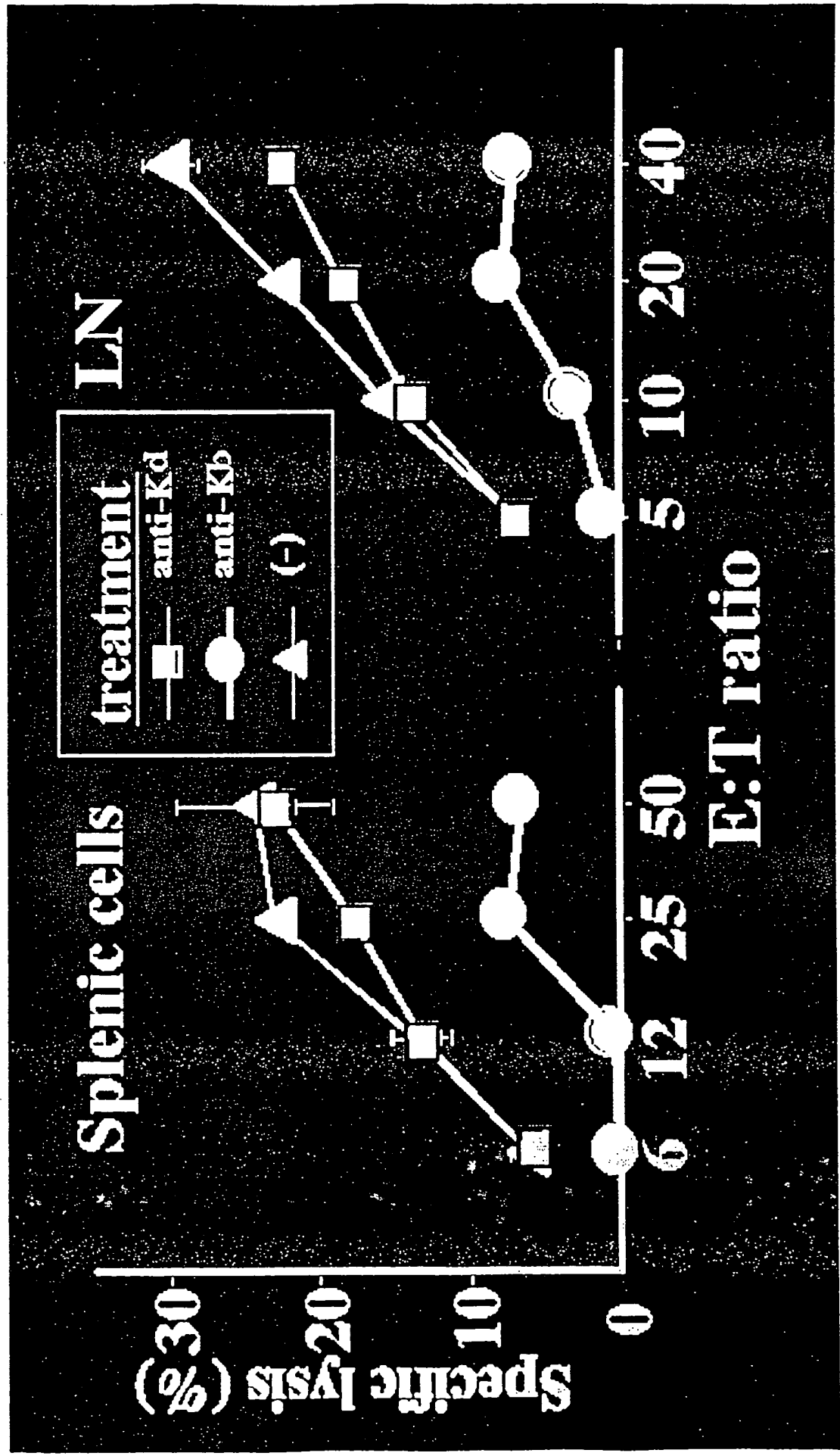


Figure 3A

Cytolytic Activity Mediated by DC and IL18

Injection is MHC Class I [K^b] Restricted



DCs Require IL12 Production to Mediate Antitumor Effects with IL18

